

inducible upstream activating sequence, 2) a minimal promoter sequence and 3) 5' and 3' P transposable elements;

(c) producing progeny from the breeding of the first *D. melanogaster* with the second *D. melanogaster*;

(d) screening the progeny for increased or decreased polyglutamine toxicity relative to the first *D. melanogaster* thereby identifying a progeny having increased or decreased polyglutamine toxicity; and

(e) identifying one or more genes operationally-associated with the marker sequence, or having an insertion of the marker sequence, that confers increased or decreased polyglutamine toxicity in the progeny having increased or decreased polyglutamine toxicity.

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10. (Twice Amended) The method of claim 1, wherein the second *D. melanogaster* is selected from a group of two or more animals having markers inserted into different locations of its genomic DNA.

11. (Twice Amended) The method of claim 10, wherein the second *D. melanogaster* is selected from a group of 10 to 100, 100 to 500, or 500 or more of the animals.

12. (Twice Amended) The method of claim 1, wherein the second *D. melanogaster* is selected from a library of animals having markers inserted at random locations of their genomic DNA.

13. (Twice Amended) The method of claim 12, wherein the library is generated by random P element insertion.

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25. (Twice Amended) A progeny *D. melanogaster* produced by the method of claim 1.

B<sup>3</sup>  
26. (Twice Amended) A transgenic *D. melanogaster* comprising a transgene containing a plurality of CAG's and at least one CAA sequence encoding a polyglutamine repeat sequence, wherein the repeat comprises at least 100 contiguous glutamine residues.

B<sup>4</sup>  
29. (Twice Amended) The *D. melanogaster* of claim 26, wherein the number of CAG's to CAA's is in ratio of between about 1:1 and 2:1.

30. (Twice Amended) The *D. melanogaster* of claim 26, wherein the number of CAG's to CAA's is in ratio of between about 2:1 and 5:1.

31. (Twice Amended) The *D. melanogaster* of claim 26, wherein the number of CAG's to CAA's is in ratio of between about 5:1 and 10:1.

32. (Twice Amended) The *D. melanogaster* of claim 26, wherein the number of CAG's to CAA's is in ratio of between about 10:1 and 50:1.

33. (Twice Amended) The *D. melanogaster* of claim 26, wherein expression of the polyglutamine sequence is conferred by a constitutive, regulatable or tissue specific expression control element.

34. (Twice Amended) The *D. melanogaster* of claim 33, wherein the tissue specific expression control element confers neural, retinal, muscle or mesoderm cell expression.

35. (Twice Amended) The *D. melanogaster* of claim 33, wherein the tissue specific expression control element comprises an Appl or rhodopsin 1 promoter or GLASS transcription factor element.

By 36. (Twice Amended) The *D. melanogaster* of claim 26, wherein the polyglutamine sequence is between about 30 and 50 amino acids in length.

37. (Twice Amended) The *D. melanogaster* of claim 26, wherein the polyglutamine sequence is between about 50 and 100 amino acids in length.

38. (Twice Amended) The *D. melanogaster* of claim 26, wherein the polyglutamine sequence is between about 100 and 200 amino acids in length.

39. (Twice Amended) The *D. melanogaster* of claim 26, wherein the polyglutamine sequence is between about 50 and 200 amino acids in length.

40. (Twice Amended) The *D. melanogaster* of claim 26, wherein the polyglutamine sequence further comprises a tag.

41. (Twice Amended) The *D. melanogaster* of claim 26, wherein polyglutamine toxicity is produced in one or more tissue or organs of the animal.

34 42. (Twice Amended) The *D. melanogaster* of claim 26, wherein the *Drosophila* further comprises a marker sequence inserted into its genomic DNA, wherein the marker is located adjacent to a gene or inserted into a gene whose expression or activity increases or decreases polyglutamine toxicity in the animal, and wherein the marker sequence comprises an inducible upstream activating sequence, a minimal promoter sequence and 5' and 3' P transposon elements containing terminal inverted repeats.

43. (Twice Amended) The *D. melanogaster* of claim 42, wherein the marker sequence is near or inserted into a gene containing a J domain.

44. (Twice Amended) The *D. melanogaster* of claim 43, wherein the gene is HDJ1.

45. (Twice Amended) The *D. melanogaster* of claim 43, wherein the gene is TPR2.

46. (Twice Amended) The *D. melanogaster* of claim 43, wherein the marker sequence is near an MLF gene.

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35 50. (Twice Amended) A method of producing a transgenic *D. melanogaster* characterized by suppressed polyglutamine toxicity comprising:

(a) transforming a *D. melanogaster* embryo or fertilized egg with a transgene comprising a plurality of CAA and CAG sequences encoding a polyglutamine sequence comprising at least 100 contiguous glutamine residues; and

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(b) selecting a *D. melanogaster* that exhibits  
polyglutamine toxicity.

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